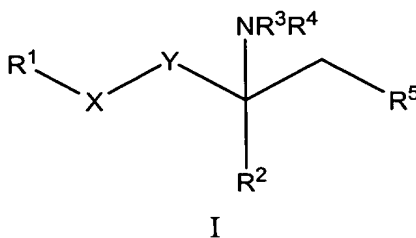


WE CLAIM

1. A compound of Formula I:

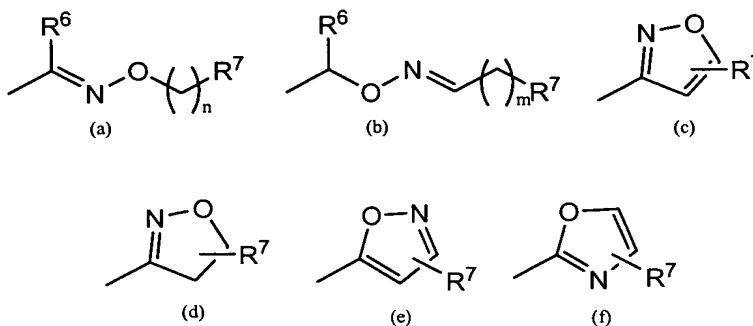


wherein:

Y is $-\text{CH}_2\text{CH}_2-$, $-\text{CH}_2\text{CH}(\text{OH})-$, $-\text{CH}(\text{OH})\text{CH}_2-$, $-\text{C}(\text{O})\text{CH}_2-$, $-\text{CH}_2\text{C}(\text{O})-$, $-\text{CH}=\text{CH}-$ or 1,2-cyclopropylene;

X is arylene or C_{5-6} heteroarylene optionally substituted by one to three substituents selected from halo, C_{1-10} alkyl and halo-substituted C_{1-6} alkyl;

R^1 is a group of formula (a), (b), (c), (d), (e) or (f):

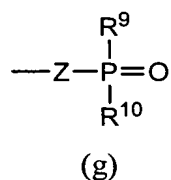


in which n is 0, 1, 2, 3, 4 or 5; m is 0, 1 or 2;

R^6 is C_{1-10} alkyl, cycloalkyl, C_{1-10} alkoxy, C_{2-10} alkenyl, C_{2-10} alkynyl, C_{1-10} alkylthio, C_{1-10} alkylsulfonyl, C_{1-10} alkylsulfinyl or halo-substituted- C_{1-10} alkyl; in each of which any aliphatic part of the group can be straight chain or branched and can be optionally substituted by up to three hydroxy, cycloalkyl, or C_{1-4} alkoxy groups and optionally interrupted by a double or triple bond or one or more $\text{C}(\text{O})$, NR^{12} , S, $\text{S}(\text{O})$, $\text{S}(\text{O})_2$ or O groups,

R⁷ is aryl or C₅₋₆heteroaryl optionally substituted by aryl, C₅₋₆heteroaryl or C₃₋₈heterocycloalkyl wherein any aryl, heteroaryl or heterocycloalkyl group of R⁷ can be optionally substituted by one to three substituents selected from halogen, C₁₋₁₀alkyl, C₁₋₁₀alkoxy, halo-substituted-C₁₋₁₀alkyl and halo-substituted-C₁₋₁₀alkoxy;

- 5 R² is hydrogen; C₁₋₄alkyl optionally substituted with one or more halogens; C₂₋₆alkenyl, C₂₋₆alkynyl, or cycloalkyl optionally substituted by halogen; or C₁₋₄alkyl optionally substituted on the terminal C atom by OH or a residue of formula (g):



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in which Z is O, S, (CH₂)₁₋₂, CF₂ or NR¹¹ where R¹¹ is H, (C₁₋₄)alkyl or halo substituted (C₁₋₄)alkyl; and R⁹ and R¹⁰, independently, are H, OH, (C₁₋₄)alkyl optionally substituted by one to three halo groups, or (C₁₋₄)alkoxy; with the proviso that R⁹ and R¹⁰ are not both hydrogen;

- 15 R³ and R⁴, independently, are H or C₁₋₄alkyl optionally substituted by halogen or acyl; and R⁵ is -OH, -Oacyl, -NHacyl, or a residue of formula (g) as defined above; or a salt thereof.

2. A compound of claim 1 wherein Y is -CH₂-CH₂- or -CH(OH)-CH₂-.

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3. A compound of claim 1 in which X is thiophenylene or phenylene.

4. A compound of claim 1 in which R¹ is a group of formula (a); R⁶ is C₁₋₆alkyl, R⁷ is thiophenyl, furanyl, pyridinyl or phenyl optionally substituted by thiophenyl, furanyl, pyridinyl, phenyl or cyclohexyl wherein any thiophenyl, furanyl, pyridinyl, phenyl or cyclohexyl of R⁷ can be optionally substituted by one to three substituents selected from halogen, halo-substituted-C₁₋₁₀alkyl and halo-substituted-C₁₋₁₀alkoxy.
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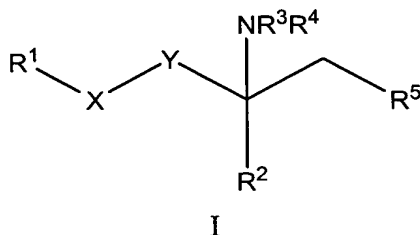
5. A pharmaceutical composition comprising a therapeutically effective amount of a compound of Claim 1 in combination with a pharmaceutically acceptable excipient.

6. A method for treating a disease in an animal in which alteration of EDG/S1P receptor mediated signal transduction can prevent, inhibit or ameliorate the pathology and/or symptomology of the disease, which method comprises administering to the animal a therapeutically effective amount of a compound of Claim 1.

7. A method for preventing or treating disorders or diseases mediated by lymphocytes, for preventing or treating acute or chronic transplant rejection or T-cell mediated inflammatory or autoimmune diseases, for inhibiting or controlling deregulated angiogenesis, or for preventing or treating diseases mediated by a neo-angiogenesis process or associated with deregulated angiogenesis in a subject comprising administering to the subject in need thereof an effective amount of a compound of claims 1, or a pharmaceutically acceptable salt thereof.

8. The use of a compound of claim 1 in the manufacture of a medicament for treating a disease in an animal in which alteration of EDG/S1P receptor mediated signal transduction contributes to the pathology and/or symptomology of the disease.

9. A process for preparing a compound of Formula I:

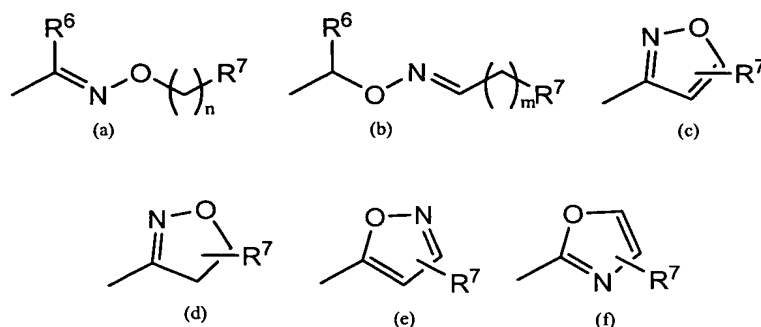


wherein:

Y is $-\text{CH}_2\text{CH}_2-$, $-\text{CH}_2\text{CH}(\text{OH})-$, $-\text{CH}(\text{OH})\text{CH}_2-$, $-\text{C}(\text{O})\text{CH}_2-$, $-\text{CH}_2\text{C}(\text{O})-$, $-\text{CH}=\text{CH}-$; or 1,2-cyclopropylene;

X is arylene or C_{5-6} heteroarylene optionally substituted by one to three substituents selected from halo, C_{1-10} alkyl and halo-substituted C_{1-6} alkyl;

5 R^1 is a group of formula (a), (b), (c), (d), (e) or (f):

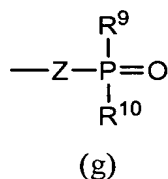


in which n is 0, 1, 2, 3, 4 or 5; m is 0, 1 or 2;

10 R^6 is C_{1-10} alkyl, cycloalkyl, C_{1-10} alkoxy, C_{2-10} alkenyl, C_{2-10} alkynyl, C_{1-10} alkylthio, C_{1-10} alkylsulfonyl, C_{1-10} alkylsulfinyl or halo-substituted- C_{1-10} alkyl; in each of which any aliphatic part of the group can be straight chain or branched and can be optionally substituted by up to three hydroxy, cycloalkyl, or C_{1-4} alkoxy groups and optionally interrupted by a double or triple bond or one or more $\text{C}(\text{O})$, NR^{12} , S, $\text{S}(\text{O})$, $\text{S}(\text{O})_2$ or O groups,

15 R^7 is aryl or C_{5-6} heteroaryl optionally substituted by aryl, C_{5-6} heteroaryl or C_{3-8} heterocycloalkyl wherein any aryl, heteroaryl or heterocycloalkyl group of R^7 can be optionally substituted by one to three substituents selected from halogen, C_{1-10} alkyl, C_{1-10} alkoxy, halo-substituted- C_{1-10} alkyl and halo-substituted- C_{1-10} alkoxy;

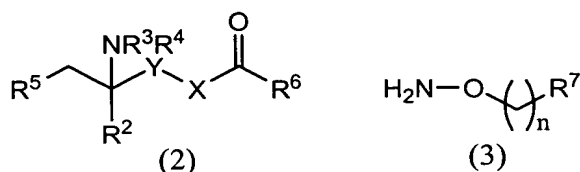
20 R^2 is hydrogen; C_{1-4} alkyl optionally substituted with one or more halogens; C_{2-6} alkenyl, C_{2-6} alkynyl, or cycloalkyl optionally substituted by halogen; or C_{1-4} alkyl optionally substituted on the terminal C atom by OH or a residue of formula (g):



in which Z is O, S, (CH₂)₁₋₂, CF₂ or NR¹¹ where R¹¹ is H, (C₁₋₄)alkyl or halo substituted (C₁₋₄)alkyl; and R⁹ and R¹⁰, independently, are H, OH, (C₁₋₄)alkyl optionally substituted by one to three halo groups, or (C₁₋₄)alkoxy; with the proviso that R⁹ and R¹⁰ are not both hydrogen;

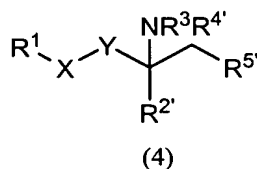
R³ and R⁴, independently, are H or C₁₋₄alkyl optionally substituted by halogen or acyl; and R⁵ is -OH, -Oacyl, -NHacyl, or a residue of formula (g) as defined above; which process comprises:

(a) reacting a compound of Formula (2) with a compound of Formula (3):



in which n, R², R³, R⁴, R⁵, R⁶, R⁷, X and Y are as defined for Formula I above; or

(b) removing the hydrolysable groups present in a compound of Formula 4:



in which R¹, R^{2'}, R³, R^{4'}, R^{5'}, X and Y are as defined for Formula I above; or

(c) optionally converting a compound of the invention into a pharmaceutically acceptable salt;

(d) optionally converting a salt form of a compound of the invention to a non-salt form;

(e) optionally converting an unoxidized form of a compound of the invention into a pharmaceutically acceptable N-oxide;

(f) optionally converting an N-oxide form of a compound of the invention to its unoxidized form;

(g) optionally resolving an individual isomer of a compound of the invention from a mixture of isomers;

(h) optionally converting a non-derivatized compound of the invention into a pharmaceutically acceptable prodrug derivative; and

(i) optionally converting a prodrug derivative of a compound of the invention to its non-derivatized form.